

Drug Utilization Criteria For Outpatient Use Guidelines Attention Deficit Disorder (ADD)/ Attention Deficit Hyperactivity Disorder (ADHD) Medications

About

Information on indications for use or diagnosis is assumed to be unavailable. All criteria may be applied retrospectively; prospective application is indicated with an asterisk [*]. The information contained is for the convenience of the public. The Texas Health and Human Services Commission is not responsible for any errors in transmission or any errors or omissions in the document.

Publication History

Revised December 2016; March 2015; February 2013; December 2012; May 2011; April 2011; March 2011; July 2008. Developed October 2007.

1. Dosage [*]

Adults

The maximum adult recommended doses for available psychostimulants used in ADD/ADHD management are summarized in Table 1.

Table 1: Maximum Recommended Adult Dosages for ADHD Medication				
		Maximum		
Drug Name	Dosage Form/Strength	Daily Dose		
STIMULANTS				
Amphetamine salts (n	nixed)*			
Adderall®, generic	tablet: 5 mg, 7.5 mg, 10 mg, 12.5 mg, 15 mg, 20 mg, 30 mg	40 mg		
Adderall XR®,	extended-release (ER) capsule: 5 mg, 10 mg, 15 mg, 20 mg,	20 mg		
generic	25 mg, 30 mg	20 1119		
Dexmethylphenidate				
Focalin®, generic	tablets: 2.5 mg, 5 mg, 10 mg	20 mg		
Focalin® XR,	ER tablets: 5mg, 10 mg, 15 mg, 20 mg, 25 mg, 30 mg, 35 mg,	40 mg		
generic	40 mg	40 1119		
Lisdexamfetamine				
Vyvanse®	capsules: 10 mg, 20 mg, 30 mg. 40 mg, 50 mg, 60 mg, 70 mg	70 mg		
Methylphenidate				
Immediate-Release:				
Methylin®, generic	chewable tablet: 2.5 mg, 5 mg, 10 mg			
Methylin®, generic	solution: 5 mg/5 mL (500 mL); 10 mg/5 mL (500 mL)	60 mg/day		
Ritalin®, generic	tablet: 5 mg, 10 mg, 20 mg			
Extended-release:				
Concerta®, generic	ER tablet: 18 mg, 27 mg, 36 mg, 54 mg	72 mg/day		
Aptensio XR®	ER capsule: 10 mg, 15 mg, 20 mg, 30 mg, 40 mg, 50 mg, 60 mg			
Metadate® CD, generic	ER capsule: 10 mg, 20 mg, 30 mg, 40 mg, 50 mg, 60 mg			
Metadate® ER,	ER tablet: 10 mg (generic only), 20 mg			
generic		60 mg/day		
Quillivant XR®	ER suspension: 300 mg/60 ml, 600 mg/120 ml, 750 mg/150 ml, 900 mg/180 ml			
QuilliChew ER™	ER chewable tablet: 20 mg, 30 mg, 40 mg			
Ritalin® LA, generic	ER capsule: 10 mg, 20 mg, 30 mg, 40 mg	1		
generic	sustained-release tablet: 20 mg			

Rev. 12/2016

File: I



Drug Utilization Criteria For Outpatient Use Guidelines
Attention Deficit Disorder (ADD)/ Attention Deficit Hyperactivity Disorder (ADHD) Medications

Table 1: Maximum Recommended Adult Dosages for ADHD Medication (continued)				
Drug Name	me Dosage Form/Strength			
NON-STIMULANTS				
SELECTIVE NOREPINEPHI	RINE REUPTAKE INHIBITORS			
Atomoxetine				
Strattera®	100 mg			

ER = extended-release

Pediatrics

Many ADHD medications are FDA-approved for use in pediatric patients. Table 2 summarizes pediatric ADD/ADHD recommended daily doses based upon patient-specific characteristics including age and weight.

Rev. 12/2016

File: I

^{*} Mixed amphetamine salts are a 1:1:1:1 combination of dextroamphetamine sulfate, dextroamphetamine saccharate, amphetamine aspartate monohydrate and amphetamine sulfate



Drug Name	Dosage Form/ Strength, Age	Maximum Daily Dose
STIMULANTS	, , , , ,	•
Amphetamine		
Evekeo®	tablet: 5 mg, 10 mg	
	• 3 -17 years of age	40 mg/day
Amphetamine salts (mix		10 mg/ day
Adderall®, generic	tablet: 5 mg, 7.5 mg, 10 mg, 12.5 mg,	
Adderant , generic	15 mg, 20 mg, 30 mg	
	• 3 -17 years of age	40 mg/day
Adderall XR®, generic	ER capsule: 5 mg, 10 mg, 15 mg, 20 mg,	+0 mg/day
Adderali XICO, generic	25 mg, 30 mg	
	• 6-12 years of age	30 mg/day
	• 13-17 years of age	20 mg/day
Daymathylabanidata	- 13-17 years or age	20 mg/day
Dexmethylphenidate	tableta: 2 F ma F ma 10	
Focalin®, generic	tablets: 2.5 mg, 5 mg, 10 mg	20/
F 1: 0 VF :	• 6-17 years of age	20 mg/day
Focalin® XR, generic	ER capsules: 5mg, 10 mg, 15 mg, 20	
	mg, 25 mg, 30 mg, 35 mg, 40 mg	
	6-17 years of age	30 mg/day
Dextroamphetamine		
Dexedrine®, generic	tablets: 5 mg, 10 mg	
	Zenzedi® capsules: 2.5 mg, 5 mg, 7,.5	
	mg, 10 mg, 15 mg, 20 mg, 30 mg;	
	ProCentra® oral solution: 5 mg/5 ml	
	3-17 years of age	40 mg/day
Dexedrine®	ER capsules: 5 mg, 10 mg, 15 mg	
Spansules, generic	 6-17 years of age 	40 mg/day ⁺
Lisdexamfetamine		
Vyvanse®	capsules: 10 mg, 20 mg, 30 mg. 40 mg,	
	50 mg, 60 mg, 70 mg	
	6-17 years of age	70 mg/day
Methamphetamine		
Desoxyn®, generic	tablet: 5 mg	
, . 5	• 6-17 years of age	25 mg/day
Methylphenidate		<u> </u>
Immediate-release		
Methylin®, generic	chewable tablet: 2.5 mg, 5 mg, 10 mg	
Methylin®, generic	solution: 5 mg/5 mL (500 mL); 10	
, 2, 32	mg/5 mL (500 mL)	
Ritalin®, generic	tablet: 5 mg, 10 mg, 20 mg	
, 3	• 6-17 years of age	60 mg/day
Extended-release :	y	23 mg/ 44 y
	FR tablet: 18 mg. 27 mg. 36 mg. 54 mg	
concertate, generic	J. J. J.	54 mg/day
	-	
Concerta®, generic	ER tablet: 18 mg, 27 mg, 36 mg, 54 mg 6-12 years 13-17 years	54 mg/day 72 mg/day (max: 2 mg/

Rev. 12/2016

File: I

Page 3 of 10 TxVendorDrug.com



Drug Utilization Criteria For Outpatient Use Guidelines Attention Deficit Disorder (ADD)/ Attention Deficit Hyperactivity Disorder (ADHD) Medications

Table 2: Pediatric (Child and Adolescent) Dosages for ADHD Medications (continued)				
Drug Name	Maximum Daily Dose			
STIMULANTS (continued)				
Methylphenidate (continued)				
Extended-release (continued)				
Aptensio XR®	ER capsule: 10 mg, 15 mg, 20 mg, 30 mg,			
	40 mg, 50 mg, 60 mg			
Metadate® CD, generic	ER capsule: 10 mg, 20 mg, 30 mg, 40 mg,			
	50 mg, 60 mg			
Metadate® ER, generic	ER tablet: 10 mg (generic only), 20 mg			
Quillivant™ XR	ER oral suspension: 300 mg/60 ml, 600			
	mg/120 ml, 750 mg/150 ml, 900 mg/180			
	ml			
QuilliChew ER™	ER chewable tablet: 20 mg, 30 mg, 40 mg			
Ritalin® LA, generic	ER capsule: 10 mg, 20 mg, 30 mg, 40 mg			
generic	sustained-release tablet: 20 mg	60 mg/day		
	6-17 years of age			
Transdermal system				
Daytrana™	transdermal patch 10 mg/9h, 15 mg/9h,			
	20 mg/9h, 30 mg/ 9h			
	6-17 years of age	30 mg/9 h/day		
NON-STIMULANTS				
SELECTIVE NOREPINEPHRINE REUP	TAKE INHIBITORS			
Atomoxetine				
Strattera®	capsules: 10 mg, 18 mg, 25 mg, 40 mg,			
	60 mg, 80 mg 100 mg			
	 6-17 years of age (≤ 70 kg) 	1.4 mg/kg/day (up		
		to 100 mg/day)		
	6-17 years of age (> 70 kg)	100 mg/day		
SELECTIVE ALPHA _{2A} -ADRENERGIC				
RECEPTOR AGONISTS				
Guanfacine				
Intuniv®, generic	ER tablet: 1 mg, 2 mg, 3 mg, 4 mg			
	• 6-12 years of age	4 mg/day		
	 13-17 years of age 	7 mg/day		

ER = extended-release

2. Duration of Therapy

Attention-deficit/hyperactivity disorder (ADHD) is defined in DSM-IV as a behavioral disorder of childhood onset characterized by symptoms of inattentiveness and hyperactivity-impulsivity. While many of the approved medications improve inattention, hyperactivity, and impulsivity in up to 70-96% of patients, symptoms may persist lifelong with less pronounced hyperactivity. Therefore, treatment often lasts well into adulthood, and ADHD is considered a chronic disorder.

3. Duplicative Therapy [*]

The use of two or more psychostimulants concurrently for ADD/ADHD management is not justified. Additional therapeutic benefit is not realized when ADHD medications are used in combination.

 Rev. 12/2016
 Page 4 of 10

 File: I
 TxVendorDrug.com

^{*} Mixed amphetamine salts are a 1:1:1:1 combination of dextroamphetamine sulfate, dextroamphetamine saccharate, amphetamine aspartate monohydrate and amphetamine sulfate

^{*}may increase dose to 60 mg/day in children weighing > 50 kg



Drug Utilization Criteria For Outpatient Use Guidelines Attention Deficit Disorder (ADD)/ Attention Deficit Hyperactivity Disorder (ADHD) Medications

Additionally, guanfacine extended-release tablets should not be prescribed concurrently with other guanfacine-containing products due to increased risks of additive pharmacologic/adverse effects, including hypotension. Patient profiles documenting receipt of multiple ADHD medications or multiple guanfacine-containing products will be reviewed.

4. Drug-Drug Interactions [*]

Patient profiles will be assessed to identify those drug regimens which may result in clinically significant drug-drug interactions. Drug-drug interactions considered clinically significant for ADD/ADHD medications are summarized in **Table 3**. Only those drug-drug interactions identified as clinical significance level 1 or those considered life-threatening which have not yet been classified will be reviewed:

Table 3: Drug-Drug Interactions for ADD/ADHD Medications				
Target Drug	Interacting Drug	Interaction	Recommendation	Clinical Significance*
amphetamines, amphetamine-related compounds, dexmethylphenidate, methylphenidate amphetamines, amphetamine-related compounds, dexmethylphenidate, methylphenidate	antihypertensive agents monoamine oxidase inhibitors (MAOIs) and drugs with MAOI-like actions (e.g., procarbazine)	combined administration decreases hypotensive effect of antihypertensive agents combined administration increases risk of enhanced vasopressor effects and hypertensive crisis due to increased norepinephrine availability; amphetamines, dexmethylphenidate, and methylphenidate potentiate catecholamine neurotransmitter effects, while MAOIS block catecholamine degradation and increase norepinephrine	closely monitor blood pressure and adjust antihypertensive therapy doses as necessary concurrent administration as well as amphetamine, dexmethylphenidate, or methylphenidate administration within 14 days of MAOI use is contraindicated	2- major (CP) contraindicated (DrugReax) 1-severe (CP)
amphetamines, amphetamine-related	phenothiazines	levels at nerve receptor sites co-administration results in decreased	avoid combination, if possible	2-major (CP) major
compounds		effectiveness of both drug classes		(DrugReax)

Rev. 12/2016

File: I

Page **5** of **10** TxVendorDrug.com



Table 3: Drug-Drug Interactions for ADD/ADHD Medications (continued)				
Target Drug	Interacting Drug	Interaction	Recommendation	Clinical Significance*
amphetamines, amphetamine- related compounds	SSRIs, SNRIs	combined administration may produce additive pharmacologic effects and increase risk of serotonin syndrome as amphetamines may stimulate serotonin release in central nervous system (CNS)	administer cautiously together and observe for signs/symptoms of serotonin syndrome; discontinue therapy and treat as necessary if serotonin syndrome develops	major (DrugReax) 2-major (CP)
amphetamines, amphetamine- related compounds	TCAs	adjunctive administration may potentiate amphetamine pharmacologic/adverse effects including hypertension, other cardiac effects, and CNS stimulation due to additive effects on norepinephrine release/activity	administer combination cautiously; observe for increased adverse effects	moderate (DrugReax) 2-major (CP)
amphetamines, amphetamine- related compounds	urinary alkalinizers	combination results in increased renal tubular absorption of amphetamines and amphetamine-related compounds, decreased urinary excretion and the potential for enhanced amphetamine therapeutic/ adverse effects	combination should be avoided	2-major (CP) moderate (DrugReax)
atomoxetine	albuterol	combined administration may produce increased heart rate, blood pressure due to unknown mechanism	administer combination cautiously; monitor blood pressure and heart rate	major (DrugReax) 3-moderate (CP)
atomoxetine	MAOIs	co-administration may result in additive serotonergic effects/increased risk of serotonin syndrome as atomoxetine inhibits serotonin reuptake and MAOIs inhibit catecholamine breakdown	concomitant administration as well as atomoxetine administration within 14 days of MAOI use contraindicated	contraindicated (DrugReax) 1-severe (CP)

Rev. 12/2016

File: I

Page 6 of 10 TxVendorDrug.com



Table 3: Drug-Drug Interactions for ADD/ADHD Medications (continued)				
Target Drug	Interacting Drug	Interaction	Recommendation	Clinical Significance*+
dexmethylphenidate, methylphenidate	select anticonvulsants [e.g., phenobarbital, hydantoins (e.g., phenytoin) and primidone]	adjunctive administration may increase serum anticonvulsant levels of select anticonvulsants due to unknown mechanism; dexmethylphenidate, methylphenidate may also lower seizure threshold	monitor serum anticonvulsant levels closely and monitor patients for increased adverse effects; adjust anticonvulsant doses as needed; also monitor seizure frequency	moderate (DrugReax) 2-major, 3- moderate (CP)
dexmethylphenidate, methylphenidate	warfarin	co-administration may increase warfarin serum levels and enhance pharmacologic/adverse effects, including bleeding	closely monitor INR with combined therapy and adjust warfarin doses as necessary	moderate (DrugReax) 3-moderate (CP)
guanfacine	antihypertensive agents	combined administration may result in additive hypotensive effects	closely monitor blood pressure and adjust doses as necessary	3-moderate (CP)
guanfacine	CNS depressants	combined administration may result in additive pharmacologic (sedative) effects	administer cautiously together	3-moderate (CP)
guanfacine	strong CYP3A4 inhibitors (e.g., ketoconazole)	adjunctive administration may result in increased guanfacine concentrations and the potential for enhanced pharmacologic/adverse effects as guanfacine is metabolized by CYP3A4	administer cautiously together and monitor for increased pharmacologic effects (e.g., hypotension, bradycardia, sedation)	unknown
guanfacine	CYP3A4 inducers (e.g., rifampin, phenytoin)	concurrent administration reduces guanfacine AUC by 70% and may result in decreased guanfacine serum levels and reduced pharmacologic/clinical effects (guanfacine metabolized by CYP3A4)	monitor for loss of guanfacine clinical effects; increased guanfacine doses may be necessary	3-moderate (CP)

Rev. 12/2016

File: I

Page 7 of 10 TxVendorDrug.com



Table 3: Drug-Drug Interactions for ADD/ADHD Medications (continued)				
Target Drug	Interacting	Interaction	Recommendation	Clinical
	Drug			Significance*
guanfacine	valproic acid	combined	monitor for additive	unknown
	(VA)	administration may	CNS effects; VA	
		result in increased VA	dosage adjustments	
		serum levels,	may be required	
		potentially due to		
		competition for		
		glucuronidation		
		metabolic pathway		
methylphenidate	bupropion	concurrent use may	if combination is	major
		result in increased	necessary, closely	(DrugReax)
mothylphopidate	carbamazonina	seizure risk	monitor patient	2-major (CP) moderate
methylphenidate	carbamazepine	co-administration may result in reduced	closely monitor patient response to	(DrugReax)
		methylphenidate serum	methylphenidate	2-major (CP)
		levels and decreased	therapy, monitor	2 major (Cr)
		pharmacologic effects	seizure frequency,	
		due to unknown	and adjust	
		mechanism;	methylphenidate	
		methylphenidate may	doses as necessary	
		also lower seizure	with this drug	
		threshold	combination	

*CP = Clinical Pharmacology

Rev. 12/2016

File: I



Drug Utilization Criteria For Outpatient Use Guidelines Attention Deficit Disorder (ADD)/ Attention Deficit Hyperactivity Disorder (ADHD) Medications

REFERENCES

- 1. Methylphenidate oral solution (Methylin®) package insert. Shionogi Pharma, Inc., **February 2015.**
- 2. Methylphenidate chewable tablets (Methylin®) package insert. Shionogi Pharma, Inc., April 2013.
- 3. Methylphenidate extended-release tablets package insert. Mallinckrodt Inc., February 2014.
- 4. Methylphenidate extended-release capsules (Metadate CD®) package insert. UCB, Inc., **February 2015**.
- 5. Methylphenidate tablets, sustained-release tablets (Ritalin®, Ritalin-SR®) package insert. Novartis Pharmaceuticals Corp., December 2013.
- 6. Methylphenidate extended-release capsules (Ritalin LA®) package insert. Novartis Pharmaceuticals Corp., **July 2015**.
- 7. Methylphenidate extended-release oral suspension (Quillivant™ XR) package insert. Pfizer, **May 2016.**
- 8. Methylphenidate extended-release chewable tablets (QuilliChew XR™) package insert. Pfizer, February 2016.
- 9. Methylphenidate extended-release capsules (Aptensio XR®) package insert. Rhodes Pharmaceuticals, May 2015.
- 10. Lisdexamfetamine (Vyvanse®) package insert. Shire US Inc., October 2016.
- 11. Dextroamphetamine sustained-release capsules (Dexedrine® Spansules®) package insert. Catalent Pharma Solutions, February 2015.
- 12. Dextroamphetamine tablets (Zenzedi®) package insert. Arbor Pharmaceuticals, LLC, June 2014.
- 13. Methamphetamine (Desoxyn®) package insert. Recordati Rare Diseases, Inc., February 2015.
- 14. Dextroamphetamine oral solution (ProCentra®) package insert. Independence Pharmaceuticals, **September 2015.**
- 15. Amphetamine mixed salts extended-release (Adderall XR®) package insert. Shire US Inc., **December 2015.**
- 16. Dextroamphetamine saccharate, amphetamine aspartate, dextroamphetamine sulfate and amphetamine sulfate tablets package insert. **Actavis Pharma, Inc., December 2015.**
- 17. Atomoxetine (Strattera®) package insert. Eli Lilly and Company, April 2015.
- 18. Dexmethylphenidate extended-release capsules (Focalin® XR) package insert. Novartis, **June 2015**.
- 19. Methylphenidate transdermal system (Daytrana®) package insert. Noven Pharmaceuticals, **August 2016.**
- 20. Guanfacine extended-release tablets (Intuniv®) package insert. Shire Pharmaceuticals Inc., **March 2016**.
- 21. DRUGDEX® System (electronic version). Truven Health Analytics, Greenwood Village, Colorado, USA. Available at: http://www.micromedexsolutions.com.libproxy.uthscsa.edu. **Accessed December 7**th, **2016**.
- 22. Clinical Pharmacology [database online]. Tampa, FL: Gold Standard, Inc.; 2016. Available at: http://www.clinicalpharmacology.com. Accessed December 7th, 2016.
- 23. Facts and Comparisons eAnswers [database online]. Hudson, Ohio: Wolters Kluwer Clinical Drug Information, Inc.; 2016. Available at: http://eanswers.factsandcomparisons.com.ezproxy.lib.utexas.edu/. Accessed December 7th, 2016.
- 24. Attention deficit/hyperactivity disorder. In: American Psychiatric Association. Diagnostic and Statistical Manual of Mental Disorders, 5th ed. Washington DC: American Psychiatric Association; 2013:59-66.
- 25. American Academy of Pediatrics. ADHD: clinical practice guideline for the diagnosis, evaluation, and treatment of attention-deficit/hyperactivity disorder in children and adolescents. Pediatrics. 2011;128:1007-22.

Rev. 12/2016 File: I



Drug Utilization Criteria For Outpatient Use Guidelines Attention Deficit Disorder (ADD)/ Attention Deficit Hyperactivity Disorder (ADHD) Medications

- 26. Volkow ND, Swanson JM. Adult attention deficit-hyperactivity disorder. N Engl J Med. 2013; 369:1935-44.
- 27. Greydanus DE, Pratt HD, Patel DR. Attention deficit hyperactivity disorder across the lifespan: the child, adolescent, and adult. Disease-A-Month. 2007;53:70-131.
- 28. Harpin VA. Medication options when treating children and adolescents with ADHD: interpreting the NICE guidance 2006. Arch Dis Child Educ Pract Ed. 2008;93:58-65.
- 29. Pliska S, for the AACAP Work Group on Quality Issues. Practice parameter for the assessment and treatment of children and adolescents with attention-deficit/hyperactivity disorder. J Am Acad Child Adolesc Psychiatry. 2007;46:894-921.
- 30. Moss SB, Nair R, Vallarino A, Wang S. Attention deficit/hyperactivity disorder in adults. Prim Care. 2007;34:445-73.
- 31. Pliska SR, Crismon ML, Hughes CW, et al, for the Texas Consensus Conference Panel on Pharmacotherapy of Childhood Attention Deficit Hyperactivity Disorder. The Texas Children's Medication Algorithm Project: revision of the algorithm for pharmacotherapy of attention-deficit/hyperactivity disorder. J Am Acad Child Adolesc Psychiatry. 2006;45(6):642-57.
- 32. Rader R, McCauley L, Callen EC. Current strategies in the diagnosis and treatment of childhood attention-deficit/hyperactivity disorder. Am Fam Physician. 2009;79(8):657-65.
- 33. Dopheide JA, Pliszka SR. Attention-deficit-hyperactivity disorder: an update. Pharmacotherapy. 2009;29(6):656-79.
- 34. DRUG-REAX® System (electronic version). Thomson Reuters (Healthcare) Inc., Greenwood Village, Colorado, USA. Available at: http://www.thomsonhc.com.libproxy.uthscsa.edu. Accessed December 7th, 2016.

Prepared by

- Drug Information Service, the University of Texas Health Science Center at San Antonio.
- The College of Pharmacy, the University of Texas at Austin.

Rev. 12/2016 File: I Page 10 of 10 TxVendorDrug.com